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FILE 'MEDLINE, AGRICOLA, DRUGU, JICST-EPLUS, CABA, BIOTECHNO, BIOSIS, CAPLUS, LIFESCI, BIOTECHDS, EMBASE, BIOENG, SCISEARCH' ENTERED AT 14:31:11 ON 31 OCT 2006

L1 36 S COQ10 AND MEVALONATE L2 3 S L1 AND PARACOCCUS

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L2 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:166835 CAPLUS

DOCUMENT NUMBER:

144:227501

TITLE:

Microbial mev operons, transgenic microorganisms containing mutant mev operons, and their use in

production of isoprenoids

INVENTOR(S):

Berry, Alan; Manhart, Christian; Simic, Petra

PATENT ASSIGNEE(S): DSM Ip Assets B. V., Neth. PCT Int. Appl., 47 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. -----____ -----______ WO 2006018211 A1 20060223 WO 2005-EP8702 20050811 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM EP 2004-19646 A 20040819 PRIORITY APPLN. INFO.: The present invention relates to a process for the production of isoprenoids, in particular Coenzyme Q10 (CoQ10) by microorganisms. More particularly, the present invention relates to a process for increased production of CoQ10 by microorganisms of the genus Rhodobacter, preferably R. sphaeroides, which have been transformed with one or more gene(s) of the mevalonate (mev) operon from a different microorganism, preferably of the genus Paracoccus, more preferably P. zeaxanthinifaciens, whereby the mev operon is mutated leading to an increased CoQ10 production Sequences carrying such a mutation as well as a microorganism carrying such a mutated mev operon are also included. Thus, Rhodobacter sphaeroides expressing the mev operon of Paracoccus containing a mutation in the hcs gene for hydroxymethylglutaryl CoA synthase produced significantly more CoOlO than did an untransformed control. THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 5 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 2 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN L_2 ACCESSION NUMBER: 2005:58354 CAPLUS 142:133188 DOCUMENT NUMBER: Improved production of coenzyme q-10 by recombinant TITLE: Rhodobacter sphaeroides Berry, Alan; Huembelin, Markus; Lopez-Ulibarri, Rual DSM IP Assets B.V., Neth. INVENTOR(S): PATENT ASSIGNEE(S): PCT Int. Appl., 19 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	ENT		KIND		DATE		APPLICATION NO.						DATE				
WO 2005005650					A1 20050120			WO 2004-EP7025						20040629			
	W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,
							LV,										
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
							ΤZ,										
	RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	ΜZ,	NA,	SD,	SL,	SZ,	ΤZ,	UG,	ZM,	ZW,	AM,

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AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
              SN, TD, TG
     EP 1641931
                           A1
                                  20060405
                                               EP 2004-740419
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK
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     US 2006154349
                                  20060713
                                               US 2006-563399
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PRIORITY APPLN. INFO.:
                                               EP 2003-15367
                                                                    A 20030708
                                               WO 2004-EP7025
                                                                    W 20040629
AΒ
     Improved process for the preparation of CoQ10 by fermentation of
     microorganisms of the genus Rhodobacter transformed with the
     mevalonate operon of Paracoccus zeaxanthinifaciens.
                                 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                                 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
      ANSWER 3 OF 3 BIOTECHDS COPYRIGHT 2006 THE THOMSON CORP. on STN
L2
ACCESSION NUMBER: 2005-06240 BIOTECHDS
                   Production of coenzyme Q10 (CoQ10), involves
TITLE:
                   introducing the mevalonate operon of
                   Paracoccus genus into Rhodobacter and cultivating the
                   modified Rhodobacter strain;
                      coenzyme production via modified bacterium culture for use
                      in food
AUTHOR:
                   BERRY A; HUEMBELIN M; LOPEZ-ULIBARRI R
                   DSM IP ASSETS BV
PATENT ASSIGNEE:
PATENT INFO:
                   WO 2005005650 20 Jan 2005
APPLICATION INFO: WO 2004-EP7025 29 Jun 2004
PRIORITY INFO: EP 2003-15367 8 Jul 2003; EP 2003-15367 8 Jul 2003
DOCUMENT TYPE:
                   Patent
                   English
LANGUAGE:
                   WPI: 2005-101913 [11]
OTHER SOURCE:
      2005-06240 BIOTECHDS
AN
      DERWENT ABSTRACT:
AR
      NOVELTY - A process (M1) for coenzyme Q10 (CoQ10) production,
      involves introducing a mevalonate operon of a microorganism
      belonging to the genus Paracoccus into a microorganism
      belonging to the genus Rhodobacter, and cultivating the modified
      Rhodobacter strain. The CoQ10 is allowed to accumulate in the
      culture and is then recovered.
            DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for: (1)
      a microorganism (I) of the genus Rhodobacter containing the
      mevalonate operon of a microorganism of the genus
      Paracoccus; (2) use of the mevalonate operon of a
      microorganism of the genus Paracoccus in (M1); and (3)
      increasing CoQ10 production in a microorganism of the genus
      Rhodobacter, involving introducing into a Rhodobacter strain the
      mevalonate operon of a microorganism of the genus
      Paracoccus and cultivating the transformant.
            BIOTECHNOLOGY - Preferred Microorganism: The Rhodobacter species
      used in the method is preferably Rhodobacter sphaeroides. The source of
      the mevalonate operon is preferably Paracoccus
      zeaxanthinifaciens.
            USE - (M1) is useful for producing coenzyme Q10 (CoQ10)
      (claimed). Coenzyme Q10 is useful, for example, as a nutritional
      supplement.
            ADVANTAGE - (M1) is an improved method for producing CoQ10
      , providing a high yield of CoQ10.
            EXAMPLE - Rhodobacter sphaeroides strain DSM 158 was used as the
      base host for construction of recombinant strains having improved
      production of coenzyme Q10 (CoQ10). Plasmids pBBR-K-Nde (empty
      plasmid) and pBBR-K-mev-op-up-4, plasmid containing the first 4 genes of
      the mevalonate operon were constructed as described in WO
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02/099095. Transformation of Escherichia coli S17-1 with plasmids for

cloning and subsequent transfer of plasmids from S17-1 to R. sphaeroides DSM 158 were performed by conjugation using standard procedures of Nishimura et al., Nucl. Acids Res.18, 6169, 1990; Simon et al., Bio/Technology 1983, 784-91. The transformed strains of R. sphaeroides DSM 158 were preserved by adding glycerol and freezing at -80 degrees C. R. sphaeroides strains DSM158, DSM 158/pBBR-K-Nde (empty vector control) and DSM 158/pBBR-K-mev-opR114 were grown in shake flask cultures, and CoQ10 production was determined. R. sphaeroides strains DSM158, DSM 158/pBBR-K-Nde and DSM 158/pBBR-K-mev-opR114 at 72 hours of culture produced 33.12 (SD 0.23), 29.71 (0.39) and 85.09 (3.42) mg/l of CoQ10, with specific formation (mg CoQ10 produced/mg cell dry mass) 2.04 (SD 0.01), 1.79 (0.10) and 3.80 (0.01), respectively. The results showed that the expression of the cloned mevalonate operon from P. zeaxanthinifaciens significantly improved CoQ10 production in R. sphaeroides.(19 pages)

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